SEP 1 1 1997 A

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of

Alan M. Warshawsky et al.

Serial No.: 08/893,080

Filed:

July 15, 1997

Title:

3-Mercaptoacetylamino-1,5-Substituted-

2-Oxo-Azepan Derivatives Useful as Inhibitors of Matrix Metalloproteinase

Examiner:

Art Unit:

I hereby certify that this correspondence is being deposited with the United States Postal Service as First Class Mail in an envelope addressed to Assistant Commissioner for Patents, Washington, D.C. 20231,

PATEN

Date of Deposit

Sanet Grubb

<u>INFORMATION DISCLOSURE STATEMENT</u> <u>UNDER 37 C.F.R. 1.56, 1.97 AND 1.98</u>

Assistant Commissioner for Patents Washington, D.C. 20231

Sir:

X.

Applicants submit herewith patents, publications, and other information of which they are aware, which they believe may be material, as defined in 37 C.F.R. 1.56(b), to the examination of this application and in respect of which there may be a duty to disclose in accordance with 37 C.F.R. 1.56(a). While the information referred to in this Information Disclosure Statement may be material pursuant to 37 C.F.R. 1.56(b), the filing of this Information Disclosure Statement is not intended to, pursuant to 37 C.F.R. 1.97(h), constitute an admission that any patent, publication or other information referred to is, or is considered to be, material to the patentability of this invention. Pursuant to 37 C.F.R. 1.97(g), the filing of this Information Disclosure Statement shall not be construed to mean that a search has been made or that no other material information exists.

(a) This Information Disclosure Statement is filed within the period set forth in §1.97(b) because it accompanies the new patent application submitted herewith, is filed within three months of the filing date of a national application or within three months of the date of entry of the national stage as set forth in §1.491 in an international application, or is believed to be filed before the mailing date of a first Office Action on the merits, whichever event occurs last. However, in the event that the first office action has been mailed, the Commissioner is authorized to charge any fees under 37 C.F.R. 1.17(p) or credit any overpayment to Account No. 13-2764.

(b)	This	s Information Disclosure Statement is filed after the period set forth in 37 C.F.R.
	1.97	(b), but is believed to be filed before the mailing date of a final action under §1.113
	or a	notice of allowance under §1.311, whichever occurs first.
	(1)	The undersigned attorney certifies that each item of information contained in this
		Information Disclosure Statement was cited in a communication from a foreign
		patent office in a counterpart foreign application not more than three months prior
		to the filing of this statement;
	(2)	The undersigned attorney certifies that no item of information contained in this
		Information Disclosure Statement was cited in a communication from a foreign
		patent office in a counterpart foreign application or, to the knowledge of the
		undersigned attorney after making reasonable inquiry, was known to any
		individual designated in §1.56(c) more than three months prior to the filing of this
		statement; or
	(3)	This Information Disclosure Statement is accompanied by a transmittal letter in
		which payment of the fee set forth in §1.17(p) and required by 37 C.F.R. 1.97(c) is
		authorized.

The items listed with on the attached PTO-1449 (modified) have not been previously submitted by the examiner or the applicant in related applications of this series. Therefore, a copy of the reference(s) are enclosed with this Information Disclosure Statement.

Respectfully submitted,

David M. Stemerick, Reg. No. 40,187

Attorney/Agent for Applicant

Hoechst Marion Roussel, Inc. 2110 East Galbraith Road P. O. Box 156300 Cincinnati, Ohio 45215-6300 Telephone (513) 948-7529 Telefax (513) 948-7961 (513) 948-4681

PATENT

heet 1 of 3

FORM PTO-1449	U.S. DEPARTMENT OF COMMERCE	ATTY, DOCKET NO.	SERIAL NO.					
(Modified) PE	PATENT AND TRADEMARK OFFICE	M01846A	08/893,080					
	RMATION DISCLOSURE	APPLICANT	APPLICANT					
CED 1 1 1997 A		Alan M. Warshawsky et al						
(Use se	eral sheets if necessary)	FILING DATE	GROUP					
TRADEMARK GE		July 15, 1997						

										U.S. PATEI	NT DOCUMENTS			
EXAMINER INITIALS	•			DC	CUN	MENT	NUM	IBER		DATE	NAME	CLASS	SUBCLASS	FILING DATE IF
		A1	4	6	8	1	9	6	6	7/21/87	Donald et al			
		A2	4	5	9	9	3	6	1	7/8/86	Dickens et al			
		A3	4	8	8	5	2	8	3	12/5/89	Broadhurst et al		<u> </u>	
		A4	4	7	7	1	0	3	8	9/13/88	Wolanin et al			
		A5	5	3	0	4	5	4	9	4/19/94	Broadhurst et al			
		A6	5	3	1	0	7	6	3	5/10/94	Campion et al			
		A7	4	9	3	5	4	0	4	6/19/90	Hunter et al			
		A8	5	2	5	2	5	6	0	10/12/93	Myers et al		-	
		A9	5	3	6	2	7	5	0	11/8/94	Rosini et al			
		A10	5	3	0	0	5	0	1	4/5/94	Porter et al			
1		A11	5	4	2	4	4	2	5	6/13/95	Flynn et al			
		A12	5	4	2	8	1	5	8	6/27/95	Warshawsky et al			
		A13	5	4	5	7	1	9	6	10/10/95	Warshawsky et al			
		A14	5	4	3	0	1	4	5	7/4/95	Flynn et al			
		A15	5	4	2	0	2	7	1	5/30/95	Warshawsky et al			
		A16	5	2	3	8	9	3	2	8/24/93	Flynn et al			
		A17	5	3	8	9	6	2	8	2/14/95	Flynn et al			
	一	A18	5	4	5	5	2	4	2	10/3/95	Warshawsky et al			<u> </u>
	\neg	A19	5	3	6	6	9	7	3	11/22/94	Flynn et al			
		A20	5	2	3	9	0	7	8	8/24/93	Galardy et al			
		A21	5	5	0	4	2	1	2	4/2/96	de Solmes et al			
					1						I	1		

FOREIGN PATENT DOCUMENTS **EXAMINER** TRANSLATION INITIALS COUNTRY DOCUMENT NUMBER DATE CLASS SUBCLASS YES **B1** 9 5 9 wo 3 2 8 5/18/95 **B2** 0 5 3 3 4 3 6 3/31/93 EP **B3** 0 8 9 5 9 6/10/92 EP **B4** 0 2 1 2 Еρ 6 0 10/26/94 **B5** 0 2 3 6 8 7 2 9/16/87 EP

	EXAMINER						DATE CON	SIDERED					
--	----------	--	--	--	--	--	----------	---------	--	--	--	--	--

EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

Not: Asterisk (*) It m(s) hav been pr viously cited in a r lated application(s) ither by the applicant or by th USPTO and therefor copies

not: Asterisk (*) it m(s) hav been pr viously cited in a r lated application(s) ither by the applicant or by th USPTO and therefor copies if the ref. r nce(s) ar not being submitted

Sheet

FORM PTO-1449 U.S. DEPARTM (Modified) PATENT AND							_					NO.		SERIAL NO. 08/893,080					
	11	NFORMA	TION	DISC	LOS	SURE					APPLICANT 00/0/3,000								
		STATEME									Alan M. Warshawsky et al								
	(Use	e several	shee	ts if r	nece	essar	y)			FILING D			GROUP	, 					
										July 15,	1997								
									U.S	S. PATENT D	ocu	MENTS							
EXAMINER INITIALS	•			DO	CUI	MENT	NUI	MBER		DATE		NAME	CLASS	SUBCLASS	FILING I				
· · · · · · · · · · · · · · · · · · ·								F	ORE	GN PATEN	T DO	CUMENTS		·					
EXAMINER INITIALS				DO	CUI	MENT	NUI	MBER		DATE		COUNTRY	CLASS	SUBCLASS	TRANSI YES	LATION NO			
		B6	9	6	3	6	2	2	7	11/21/96	wo								
		B7	9	7	1	9	0	7	5	5/29/97	wo								
		B8	9	7	1	7	0	8	8	5/15/97	wo								
EVALUED.		· · · · ·								OTHER DO	CUM	NTS			·				
EXAMINER INITIALS				AUTHOR, TITLE, DATE, PERTINENT PAGES, ETC.															
		C1	Mo	McCachren, S.S. <u>Arthritis Rheum</u> . 34, pp 1085-1093 (1991)															
		C2	En	Emonard, H. et al, Cell Molec. Biol. 36, pp 131-153 (1990)															
		C3	Bi	Birkedal-Hansen, H., J. Oral Pathol. 17, pp 445-451 (1988)															
		C4	Ma	Matrisian, L.M., <u>Trends Genet</u> . 6, pp 121-125 (1990)															
	1	C5	Mı	urpł	ıy,	G.J.	Ρ. ϵ	et al,	FE	BS Lett. 2	89, p	p 4-7 (1991							
,		C6	M	atris	sian	, L.	M.,	Bio	essa	<u>ys</u> 14, pp	455-4	163 (1992)				·			
		C7	Sw	vhw	artz	z, G	.K.	et al	, <u>C</u> a	ncer 73, p	p 22-	27 (1994)							
	Ì	C8	Ве	rnh	ard	, E	l. et	al,]	Proc	. Natl. Ac	ad. S	<u>ci</u> . 91, pp 4293-	4597 (1994	4)					
		C9	Zu	cke	r, S	S. et	al,	Can	cer l	Res. 53, pp	140	-146 (1993)							
		C10	Ch	iriv	i, F	R.G.	S. e	t al,	<u>Int.</u>	J. Cancer	58, p	p 460-464 (199	4)						
-		C11	Mo	ontg	gom	nery	, A.	M.P	. et	al, Cancer	Res.	54, pp 5467-54	73 (1994)						
-		C12	Ha	sty,	, K .	Α. ε	t al	, <u>Ar</u>	thr.	Rheum. 33	3, pp	388-397 (1990)				_			
		C13	Μι	urph	ıy,	G. e	t al	, <u>Bic</u>	che	<u>m. J</u> . 248,	pp 2	65-268 (1987)			_				
		C14	Ge	arin	ıg,	A.J.	H. (et al,	<u>Na</u>	ture 370, p	p 55	5-557 (1994)							
		C15	Mo	ohle	r, F	K.M	. et	al, <u>l</u>	Vatu	<u>re</u> 370, pp	218-	220 (1994)	· · · · · · · · · · · · · · · · · · ·						
		C16	Mo	Gre	eeh	an,	G.N	1. et	al,]	Nature 370), pp	558-561 (1994)							
EXAMINER											DATE	CONSIDERED							

EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. Note: Ast risk (*) it m(s) hav been pr viously cited in a related application(s) ither by th applicant r by th USPTO and ther for copi s of the r fer nce(s) ar not being submitted.

DATE CONSIDERED

Sheet

FORM PTO-1449	U.S. DEPARTMENT OF COMMERCE	ATTY, DOCKET NO.	SERIAL NO.				
(Modified)	PATENT AND TRADEMARK OFFICE	M01846A	08/893,080				
	FORMATION DISCLOSURE	APPLICANT					
ST	ATEMENT BY APPLICANT	Alan M. Warshawsky					
(Use	several sheets if necessary)	FILING DATE	GROUP				
		July 15, 1997					

FOREIGN PATENT DOCUMENTS

EXAMINER INITIALS	*			DO	CUM	1ENT	NUM	BER		DATE	COUNTRY	CLASS	SUBCLASS	TRANSLATION YES NO			
		В9	9	7	1	2	9	0	2	4/10/97	wo						
		B10	9	7	1	2	8	6	1	4/10/97	wo						
		B11	9 6 3 5 7 1 4				1	4	11/14/96	wo							
-		B12	9	6	6	5	7	1	2	11/14/96	wo						
		B13	9	6	3	5	7	1	1	11/14/96	wo	1					
		B14	9	6	3	5	6	8	7	11/14/96	wo						

OTHER DOCUMENTS

		OTHER DOCOMENTS
EXAMINER INITIALS		AUTHOR, TITLE, DATE, PERTINENT PAGES, ETC.
	C17	Wahl, R.C. et al, Annual Reports in Medicinal Chemistry 25, pp 177-184 (1990)
	C18	Hennney, A.m., et al, Proc. Natl. Acad. Sci. 88, pp 8154-8158 (1991)
	C19	Burns, F.R. et al, Invest. Opthalmol. and Visual Sci. 30, No. 7 pp 1569-1575 (1989)
	C20	Overall, C.M. et al, <u>J. Periodontal Res</u> . 22, pp 81-88 (1987)
	C21	Miyazaki, K. et al, <u>Nature</u> 362, pp 839-841 (1993)
	C22	Beckett, R.P. et al, "Recent advances in matrix metalloproteinase inhibitor research" <u>DDT</u> Vol. 1, No. 1, January, 1996, pp 16-26
	C23	Blanchard et al, "Reevaluation of the Absorption of Carbenoxolone Using an In Situ Rat Intestinal Techinque", Journ. Pharmaceutical Sci., Vol. 79, No. 5, May, 1990 pp 411-414
	C24	Davies, B. et al, "A Synthetic Matrix Metalloproteinase Inhibitor Decreases Tomor Burden", Cancer Research 53, May 1, 1993, pp 2087-2091
	C25	Hodgson, John, "Remodeling MMPIs", Bio/Technology Vol. 13, June 1995 pp 554-557
	C26	Chapman, K.T., "Inhibition of Matrix Metalloproteinases", J. Med. Chem. 1993, 36, pp 4293-4301
	C27	Beeley N.RA, "Inhibitors of Matrix Metalloproteinases (MMP's)", Curr. Opin, Ther. Patents (1994) 4(1), pp 7-16
	C28	Baxter, Andrew D. et al, "A Novel Series of Matrix Metalloproteinase Inhibitors for the Treatment of Inflammatory Disorders", Bioorganic & Medicinal Chem. Letters, Vol. 7, No. 7 pp 897-902, (1997)
	C29	CA, Issue 3, pp 16, 126:60361y (1997)

EXAMINER	DATE CONSIDERED

EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in

conformance and not considered. Include copy of this form with next communication to applicant.

Not: Ast risk (*) it m(s) hav been pr viously cited in a r lated application(s) ither by the applicant r by th USPTO and th r for copi s of the r ferenc (s) ar not being submitted.